

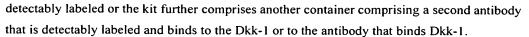
WHAT IS CLAIMED IS:

- 1. A method of treating insulin resistance or hypoinsulinemia in mammals comprising administering to a mammal in need thereof an effective amount of an antagonist to Dickkopf-1 (Dkk-1).
- The method of claim 1 wherein the mammal has non-insulin dependent diabetes mellitus (NIDDM).
- 3. The method of claim 1 wherein the mammal is human and the antagonist is to human Dkk-1.
- 4. The method of claim 1 wherein the antagonist is an antibody that binds Dkk-1.
- 5. The method of claim 4 wherein the antibody is a monoclonal antibody.
- 6. The method of claim 5 wherein the antibody is prepared from a hybridoma having ATCC Dep. No. PTA-3086.
- 7. The method of claim 1 wherein the administration is systemic.
- 8. The method of claim 1 wherein insulin resistance is treated, further comprising administering an effective amount of an insulin-resistance-treating agent to the mammal.
- 9. The method of claim 1 wherein hypoinsulinemia is treated, further comprising administering an effective amount of insulin to the mammal.
- 10. A method for detecting the presence or onset of insulin resistance or hypoinsulinemia in a mammal comprising the steps of:
 - (a) measuring the amount of Dickkopf-1 (Dkk-1) in a sample from said mammal; and
 - (b) comparing the amount determined in step (a) to an amount of Dkk-1 present in a standard sample, an increased level in the amount of Dkk-1 in step (a) being indicative of insulin resistance or hypoinsulinemia.
- 11. The method of claim 10 wherein the measuring is carried out using an anti-Dkk-1 antibody in an immunoassay.
- 12. The method of claim 11 wherein the anti-Dkk-1 antibody comprises a label.
- 13. The method of claim 12 wherein the label is selected from the group consisting of a fluorescent label, a radioactive label, and an enzyme label.
- 14. The method of claim 11, wherein the immunoassay is selected from the group consisting of a radioimmunoassay, an enzyme immunoassay, an enzyme-linked immunosorbent assay, a sandwich immunoassay, a precipitation assay, an immunoradioactive assay, a fluoresence immunoassay, a protein A immunoassay, and an immunoelectrophoresis assay.
- 15. The method of claim 10 wherein the insulin resistance is non-insulin dependent diabetes mellitus.
- 16. The method of claim 10 wherein the mammal is human and human Dkk-1 is being measured.
- 17. A kit for treating insulin resistance or hypoinsulinemia, said kit comprising:
 - (a) a container comprising an antagonist to Dkk-1; and
 - (b) instructions for using the antagonist to treat insulin resistance or hypoinsulinemia.
- 18. The kit of claim 17 wherein the antagonist is an antibody that binds Dkk-1.
- 19. The kit of claim 18 wherein the antibody is a monoclonal antibody.
- 20. The kit of claim 18 wherein the antibody binds human Dkk-1.
- 21. The kit of claim 17 for treating non-insulin dependent diabetes.





- 22. The kit of claim 17 further comprising a container comprising an insulin-resistance-treating agent if insulin resistance is treated or insulin if hypoinsulinemia is treated.
- 23. A hybridoma selected from the group consisting of ATCC Dep. No. PTA-3084, PTA-3085, PTA-3086, PTA-3087, PTA-3088, PTA-3089, and PTA-3097.
- 24. The hybridoma of claim 23 that is ATCC Dep. No. PTA-3086.
- 25. An antibody prepared from the hybridoma of claim 23.
- 26. A method of treating obesity or hyperinsulinemia in mammals comprising administering to a mammal in need thereof an effective amount of Dickkopf-1 (Dkk-1).
- 27. The method of claim 26 wherein the mammal is human and the Dkk-1 is human Dkk-1.
- 28. The method of claim 26 wherein the administration is systemic.
- 29. The method of claim 26 further comprising administering an effective amount of weight-loss agent.
- 30. A method for detecting the presence or onset of obesity or hyperinsulinemia in a mammal comprising the steps of:
 - (a) measuring the amount of Dickkopf-1 (Dkk-1) in a sample from said mammal; and
 - (b) comparing the amount determined in step (a) to an amount of Dkk-1 present in a standard sample, a decreased level in the amount of Dkk-1 in step (a) being indicative of obesity or hyperinsulinemia.
- 31. The method of claim 30 wherein the measuring is carried out using an anti-Dkk-1 antibody in an immunoassay.
- 32. The method of claim 31 wherein the anti-Dkk-1 antibody comprises a label.
- 33. The method of claim 32 wherein the label is selected from the group consisting of a fluorescent label, a radioactive label, and an enzyme label.
- 34. The method of claim 31, wherein the immunoassay is selected from the group consisting of a radioimmunoassay, an enzyme immunoassay, an enzyme-linked immunosorbent assay, a sandwich immunoassay, a precipitation assay, an immunoradioactive assay, a fluoresence immunoassay, a protein A immunoassay, and an immunoelectrophoresis assay.
- 35. The method of claim 30 wherein the mammal is human and human Dkk-1 is being measured.
- 36. A kit for treating obesity or hyperinsulinemia, said kit comprising:
 - (a) a container comprising Dkk-1; and
 - (b) instructions for using the Dkk-1 to treat obesity or hyperinsulinemia.
- 37. The kit of claim 36 wherein the Dkk-1 is human Dkk-1.
- 38. The kit of claim 36 further comprising a container comprising a weight-loss agent if obesity is being treated or comprising diazoxide if hyperinsulinemia is being treated.
- 39. A diagnostic kit for detecting the presence or onset of insulin resistance, hyperinsulinemia, hypoinsulinemia, or obesity, said kit comprising:
 - (a) a container comprising an antibody that binds Dickkopf-1 (Dkk-1);
 - (b) a container comprising a standard sample containing Dkk-1; and
 - (c) instructions for using the antibody and standard sample to detect insulin resistance, hyperinsulinemia, hypoinsulinemia, or obesity, wherein either the antibody that binds Dkk-1 is



- 40. The kit of claim 39 wherein the antibody that binds Dkk-1 is a monoclonal antibody.
- 41. The kit of claim 39 wherein the Dkk-1 is human Dkk-1 and the kit is for detecting non-insulin dependent diabetes or obesity.
- 42. A method for repairing or regenerating muscle in a mammal comprising administering to the mammal an effective amount of an antagonist to Dkk-1.
- 43. The method of claim 42 wherein the antagonist is an antibody that binds Dkk-1.
- 44. The method of claim 43 wherein the mammal is human and the antibody binds human Dkk-1.
- 45. The method of claim 42 wherein the antibody is a monoclonal antibody.
- 46. A kit for repairing or regeneration muscle, said kit comprising:
 - (a) a container comprising an antagonist to Dkk-1; and
 - (b) instructions for using the antagonist to repair or regenerate muscle in a mammal.
- 47. A monoclonal antibody preparation prepared by hyperimmunizing mice with tagged Dkk-1 diluted in an adjuvant, fusing B-cells from the mice having anti-Dkk-1 antibody titers with mouse myeloma cells and obtaining supernatants, harvesting the supernatants, screening the harvested supernatants for antibody production, injecting positive clones showing the highest immunobinding after a second round of subcloning into primed mice for *in vivo* production of monoclonal antibodies, pooling ascites fluids from the mice, and purifying the ascites fluid pool to produce the antibody preparation.
- 48. A method of evaluating the effect of a candidate pharmaceutical drug on insulin resistance, hypoinsulinemia, or muscle repair comprising administering said drug to a non-human transgenic animal that overexpresses *dkk-1* nucleic acid and determining the effect of the drug on glucose clearance from the blood of said animal, on circulating insulin levels in said animal, or on muscle differentiation, respectively.
- 49. A method of evaluating the effect of a candidate pharmaceutical drug on obesity or hyperinsulinemia comprising administering said drug to a non-human binary transgenic animal that expresses *dkk-1* nucleic acid and determining the effect of the drug on an obesity-determining property or on the level of insulin in said animal.
- 50. A non-human transgenic animal that overexpresses dkk-1 nucleic acid.
- 51. The animal of claim 50 that is a rodent.
- 52. The animal of claim 50 that is a mouse.